

<110> Balint, Robert F. Her, Jeng-Horng KaloBios, Inc.

<120> Interaction-Activated Proteins

<130> 021167-000700US

<140> US 09/526,106

<141>2000-03-15

<150> US 60/124,339

<151> 1999-03-15

<150> US 60/135,926

<151> 1999-05-25

<150> US 60/175,968

<151>2000-01-13

<160> 26

<170> PatentIn Ver. 2.1

<210>1

<211> 789

<212> DNA

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<223> TEM-1 beta-lactamase

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gca cga gtg ggt tac atc gaa ctg gat ctc aac agc ggt aag atc ctt Ala Arg Val Gly Tyr Ile Glu Leu Asp Leu Asn Ser Gly Lys Ile Leu

96

48

20 25 30

•	
Glu Ser Phe Arg Pro Glu Glu Arg Phe Pro Met Met Ser Thr Phe Lys	
35 40 45	
gtt ctg cta tgt ggc gcg gta tta tcc cgt att gac gcc ggg caa gag Val Leu Cys Gly Ala Val Leu Ser Arg Ile Asp Ala Gly Gln Glu 50 55 60	192
caa ctc ggt cgc cgc ata cac tat tct cag aat gac ttg gtt gag tac Gln Leu Gly Arg Arg Ile His Tyr Ser Gln Asn Asp Leu Val Glu Tyr 65 70 75 80	240
tca cca gtc aca gaa aag cat ctt acg gat ggc atg aca gta aga gaa Ser Pro Val Thr Glu Lys His Leu Thr Asp Gly Met Thr Val Arg Glu 85 90 95	288
tta tgc agt gct gcc ata acc atg agt gat aac act gcg gcc aac tta Leu Cys Ser Ala Ala Ile Thr Met Ser Asp Asn Thr Ala Ala Asn Leu 100 105 110	336
ctt ctg aca acg atc gga gga ccg aag gag cta acc gct ttt ttg cac Leu Leu Thr Thr Ile Gly Gly Pro Lys Glu Leu Thr Ala Phe Leu His 115 120 125	384
aac atg ggg gat cat gta act cgc ctt gat cgt tgg gaa ccg gag ctg Asn Met Gly Asp His Val Thr Arg Leu Asp Arg Trp Glu Pro Glu Leu 130 135 140	432
aat gaa gcc ata cca aac gac gag cgt gac acc acg atg cct gta gca Asn Glu Ala Ile Pro Asn Asp Glu Arg Asp Thr Thr Met Pro Val Ala 145 150 155 160	480
atg gca aca acg ttg cgc aaa cta tta act ggc gaa cta ctt act cta Met Ala Thr Thr Leu Arg Lys Leu Leu Thr Gly Glu Leu Leu Thr Leu 165 170 175	528
get tee egg caa caa tta ata gae tgg atg gag geg gat aaa gtt gea Ala Ser Arg Gln Gln Leu Ile Asp Trp Met Glu Ala Asp Lys Val Ala 180 185 190	576
gga cca ctt ctg cgc tcg gcc ctt ccg gct ggc tgg ttt att gct gat Gly Pro Leu Leu Arg Ser Ala Leu Pro Ala Gly Trp Phe Ile Ala Asp 195 200 205	624
aaa tct gga gcc ggt gag cgt ggg tct cgc ggt atc att gca gca ctg Lys Ser Gly Ala Gly Glu Arg Gly Ser Arg Gly Ile Ile Ala Ala Leu 210 215 220	672

ggg cca gat ggt aag ccc tcc cgt atc gta gtt atc tac acg acg ggg Gly Pro Asp Gly Lys Pro Ser Arg Ile Val Val Ile Tyr Thr Thr Gly 225 230 235 240	720
agt cag gca act atg gat gaa cga aat aga cag atc gct gag ata ggt Ser Gln Ala Thr Met Asp Glu Arg Asn Arg Gln Ile Ala Glu Ile Gly 245 250 255	768
gcc tca ctg att aag cat tgg Ala Ser Leu Ile Lys His Trp 260	789
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<220> <223> TEM-1 beta-lactamase	
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Ala Arg Val Gly Tyr Ile Glu Leu Asp Leu Asn Ser Gly Lys Ile Leu 20 25 30	
Glu Ser Phe Arg Pro Glu Glu Arg Phe Pro Met Met Ser Thr Phe Lys 35 40 45	
Val Leu Cys Gly Ala Val Leu Ser Arg Ile Asp Ala Gly Gln Glu 50 55 60	
Gln Leu Gly Arg Arg Ile His Tyr Ser Gln Asn Asp Leu Val Glu Tyr 65 70 75 80	
Ser Pro Val Thr Glu Lys His Leu Thr Asp Gly Met Thr Val Arg Glu 85 90 95	
Leu Cys Ser Ala Ala Ile Thr Met Ser Asp Asn Thr Ala Ala Asn Leu 100 105 110	
Leu Leu Thr Thr Ile Gly Gly Pro Lys Glu Leu Thr Ala Phe Leu His 115 120 125	

Asn Met Gly Asp His Val Thr Arg Leu Asp Arg Trp Glu Pro Glu Leu 130 135 140 Asn Glu Ala Ile Pro Asn Asp Glu Arg Asp Thr Thr Met Pro Val Ala 160 145 150 155 Met Ala Thr Thr Leu Arg Lys Leu Leu Thr Gly Glu Leu Leu Thr Leu 175 165 170 Ala Ser Arg Gln Gln Leu Ile Asp Trp Met Glu Ala Asp Lys Val Ala 180 185 190 Gly Pro Leu Leu Arg Ser Ala Leu Pro Ala Gly Trp Phe Ile Ala Asp 195 200 205 Lys Ser Gly Ala Gly Glu Arg Gly Ser Arg Gly Ile Ile Ala Ala Leu 210 215 220 Gly Pro Asp Gly Lys Pro Ser Arg Ile Val Val Ile Tyr Thr Thr Gly 235 240 225 230 Ser Gln Ala Thr Met Asp Glu Arg Asn Arg Gln Ile Ala Glu Ile Gly 245 250 255 Ala Ser Leu Ile Lys His Trp 260 <210>3 <211>5 <212> PRT <213> Artificial Sequence <220> <223> Description of Artificial Sequence:linker <400> 3 Gly Gly Gly Ser 5 1 <210>4 <211> 15 <212> PRT <213> Artificial Sequence

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                                                     15
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<400> 5
His His His His His
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               5
<210>6
<211>5
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   of variable length
<220>
<221> REPEAT
<222>(1)..(5)
<223> (G-4S)-x, amino acids 1-5 may be repeated an
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<400>6
Gly Gly Gly Ser
                5
1
<210>7
<211>267
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<213 > Esche	erichia coli					
<220> <223> Neon	nycin phos	photransfe	erase II (N	NPTII)		
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Ala Trp Val	Glu Arg L 20	eu Phe Gl	y Tyr As <sub>j</sub> 25	p Trp Ala	Gln Glr 30	
Gly Cys Ser 35	Asp Ala A	la Val Pho 40	_	u Ser Ala	Gln Gly 45	Arg Pro
Val Leu Phe 50	Val Lys T	hr Asp Le 55	u Ser Gly	y Ala Leu 60	ı Asn Glı	ı Leu Gln
Asp Glu Ala 65	_	eu Ser Trj 70	p Leu Ala	a Thr Thr 75	Gly Val	Pro Cys 80
Ala Ala Val	Leu Asp V 85	al Val Th	r Glu Ala 90		Asp Trp	Leu Leu 95
Leu Gly Glu	Val Pro G 100	ly Gln As	p Leu Le 105	u Ser Ser	His Leu 110	
Ala Glu Lys	Val Ser Ile 115	e Met Ala	Asp Ala 120	Met Arg	Arg Leu 125	
Leu Asp Pro		ys Pro Ph	-	s Gln Ala	Lys His 140	Arg Ile
Glu Arg Ala 145	Arg Thr A	rg Met Gi 150	lu Ala Gl	y Leu Va 15	_	n Asp Asp
Leu Asp Glu 160		iln Gly Le 65	u Ala Pro	o Ala Glu 170	Leu Pho	e Ala Arg 175
Leu Lys Ala	Arg Met P 180	ro Asp G	ly Glu As 18:		al Val Th	r His Gly 190
Asp Ala Cys	Leu Pro A	sn Ile Me	t Val Glu 200	Asn Gly	Arg Pho	•
Phe Ile Asp	Cys Gly Aı	g Leu Gly	y Val Ala	Asp Arg	Tyr Gln	Asp Ile

220

Ala Leu Ala Thr Arg Asp Ile Ala Glu Glu Leu Gly Gly Glu Trp Ala 225 230 Asp Arg Phe Leu Val Leu Tyr Gly Ile Ala Ala Pro Asp Ser Gln Arg 240 245 250 255 Ile Ala Phe Tyr Arg Leu Leu Asp Glu Phe Phe 260 265 <210>8 <211>18 <212> PRT <213> Artificial Sequence <220> <223> Description of Artificial Sequence: CD40-binding Trxpep <400>8 Cys Gly Pro Lys Glu Leu Arg Ile Gly Gly Arg Pro Arg Pro Gly 5 10 15 Pro Cys <210>9 <211>18 <212> PRT <213> Artificial Sequence <220> <223> Description of Artificial Sequence: CD40-binding Trxpep <400>9 Cys Gly Pro Glu Gly Gln Gly Gly Val Ala Val Gly Gly Val Gly Gly 5 10 15 Pro Cys <210>10 <211>16

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                                                        15
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<210>11
<211>21
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<400>11
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                5
                                  10
                                                      15
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Leu Gln Pro Gly Ala
            20
<210> 12
<211>18
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Cys Gly Pro Lys Ser Ala Gly Lys Gly Arg Lys Asp Arg Arg Lys Gly
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1
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Pro Cys
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<210> 13

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Gly Pro Cys
<210> 14
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Cys Gly Pro Ala Gly Ala Ile Arg His Glu His Arg Gln Gly Leu Gly
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Pro Cys
<210> 15
<211>23
<212> PRT
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   Trxpep
<400>15
Leu Val Thr Leu Glu Asn Gly Lys Gln Leu Thr Val Lys Arg Gln Gly
                  5
                                     10
                                                         15
Leu Tyr Tyr Ile Tyr Ala Gln
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20

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<210> 16
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Cys Gly Pro Asp Thr Gly Leu Glu Thr Asp Ala Ala Asp Ala Ser Gly
                5
                                                        15
Pro Cys
<210>17
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   Trxpep
<400> 17
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                                     10
Pro Cys
<210> 18
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<212> PRT
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   Trxpep
<400> 18
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Pro Cys
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1
                5
                                   10
                                                        15
Pro Gln
<210>20
<211>18
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   Trxpep
<400> 20
Cys Gly Pro Gly Arg Glu Ser Arg Gly Arg Cys Tyr Thr Pro Ser Gly
                 5
1
                                                       15
Pro Cys
<210>21
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Thr Asp Pro Ser Gln Val Ser His Gly Thr Gly Phe Thr Ser Phe Gly
                                   10
                                                      15
Leu Leu
<210>22
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   Trxpep
<400> 22
Cys Gly Pro Asn Thr Pro Asp Glu Glu Met Ala Pro Gln Ala Pro Gly
                 5
                                    10
1
                                                       15
Pro Cys
<210> 23
<211>18
<212> PRT
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   Trxpep
<400> 23
Cys Gly Pro Val Val His Ile Lys Thr Asn Glu Gln Ala Ala Pro Gly
                                  10
Pro Cys
<210>24
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<212> PRT
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<400> 24
Cys Gly Pro Val Ala Glu Glu Pro Ala Gly Gly Ala Gly Arg Pro Gly
                                                        15
                                    10
Pro Cys
<210> 25
<211>9
<212> PRT
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   Tyr1068 phosphorylation substrate peptide
<400> 25
Pro Val Pro Glu Tyr Ile Asn Gln Ser
                5
<210>26
<211>5
<212> PRT
<213> Artificial Sequence
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<223> Description of Artificial Sequence:short flexible
   linker
<400> 26
Pro Gly Ser Gly Gly
1
                5
<210> 27
<211>263
<212> PRT
<213> Escherichia coli
<220>
<223> N-terminal beta-lactamase fragment
<400> 2
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Ala Arg Val Gly 20	Tyr Ile Glu Le	u Asp Leu Asr 25	n Ser Gly Lys 30	Ile Leu
Glu Ser Phe Arg		rg Phe Pro Me 0	t Met Ser Thr 45	Phe Lys
Val Leu Leu Cys 50	s Gly Ala Val L 55	eu Ser Arg Ile	Asp Ala Gly 60	Gln Glu
Gln Leu Gly Arg	g Arg Ile His Ty 70	yr Ser Gln Asn 75	Asp Leu Val	Glu Tyr 80
Ser Pro Val Thr	Glu Lys His Le 85	eu Thr Asp Gly 90	Met Thr Val	Arg Glu 95
Leu Cys Ser Ala 100	Ala Ile Thr Me	et Ser Asp Asn 105	Thr Ala Ala A	Asn Leu
Leu Leu Thr Thr 115	: Ile Gly Gly Pro 120	•	Thr Ala Phe 1 125	Leu His
Asn Met Gly As 130	p His Val Thr A 135	Arg Leu Asp A	rg Trp Glu Pro 140	o Glu Leu
Asn Glu Ala Ile 145	Pro Asn Asp G 150	lu Arg Asp Th 155	r Thr Thr Pro	Val Ala 160
Met Ala Thr Thr	Leu Arg Lys L 165	eu Leu Thr Gl 170	y Glu Leu Le	u Thr Leu 175
Ala Ser Arg Gln 180	Gln Leu Ile As	p Trp Met Glu 185	Ala Asp Lys 190	Val Ala
Gly Pro Leu Leu 195	Arg Ser Ala L	•	Trp Phe Ile A	Ala Asp
Lys Ser Gly Ala 210	Gly Glu Arg G 215	ly Ser Arg Gly	' Ile Ile Ala A 220	la Leu
Gly Pro Asp Gly 225	Lys Pro Ser Ar 230	rg Ile Val Val 1 235	lle Tyr Thr Th	or Gly 240
Ser Gln Ala Thr	Met Asp Glu A	arg Asn Arg Gl	n Ile Ala Glu	Ile Gly

245 250 255

Ala Ser Leu Ile Lys His Trp 260

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